

'HERMALLY STABLE AT LOWER HEATING TEMPERATURES (~50°C) **TRANSFORMATION MAY PROCEED THROUGH ANOTHER TERM** 

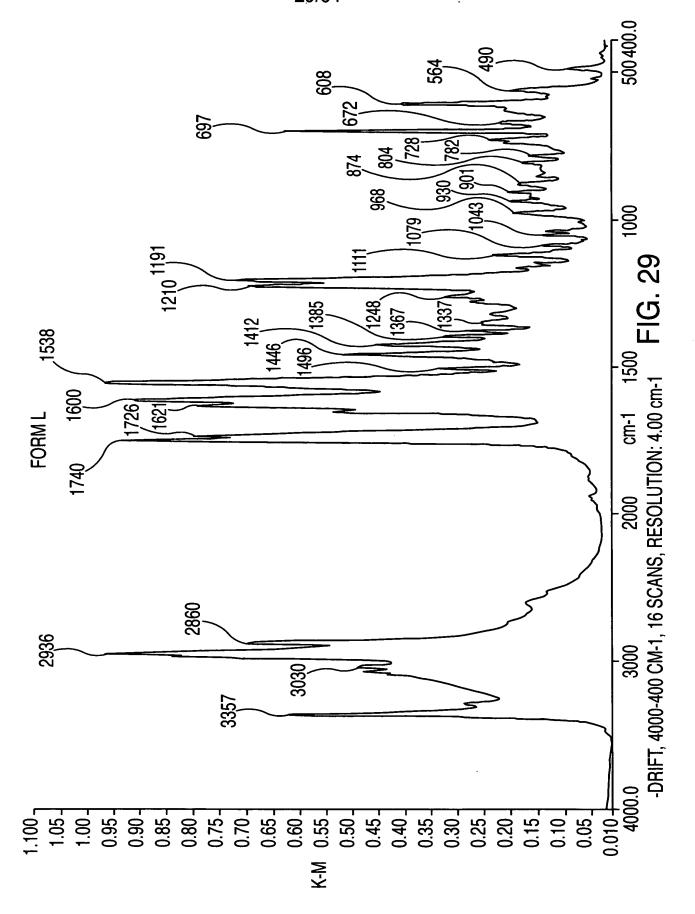
HERMALLY STABLE FORMS.

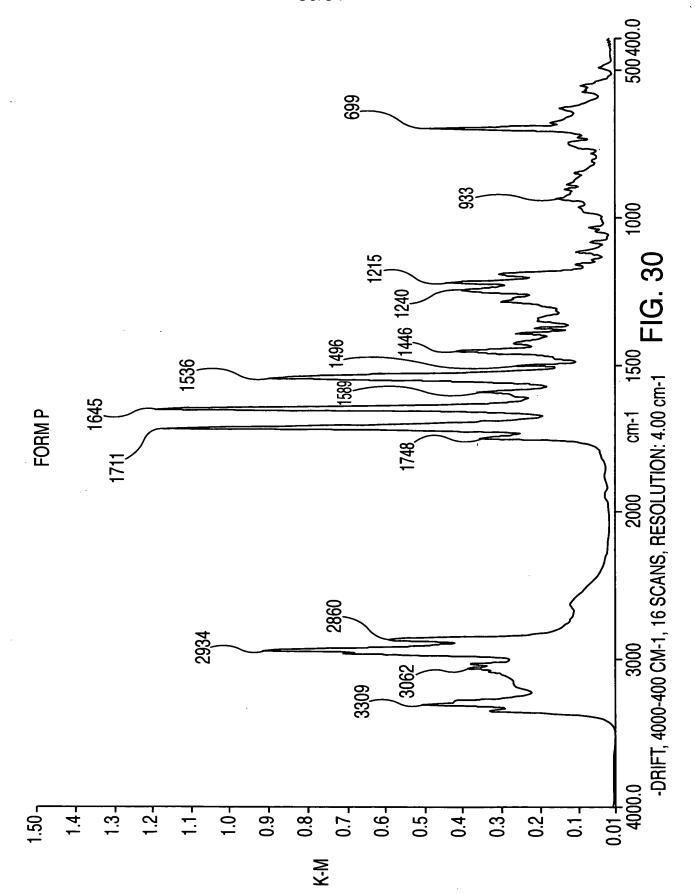
RANSFORMATION AFTER STORAGE AT ROOM TEMPERATURE.

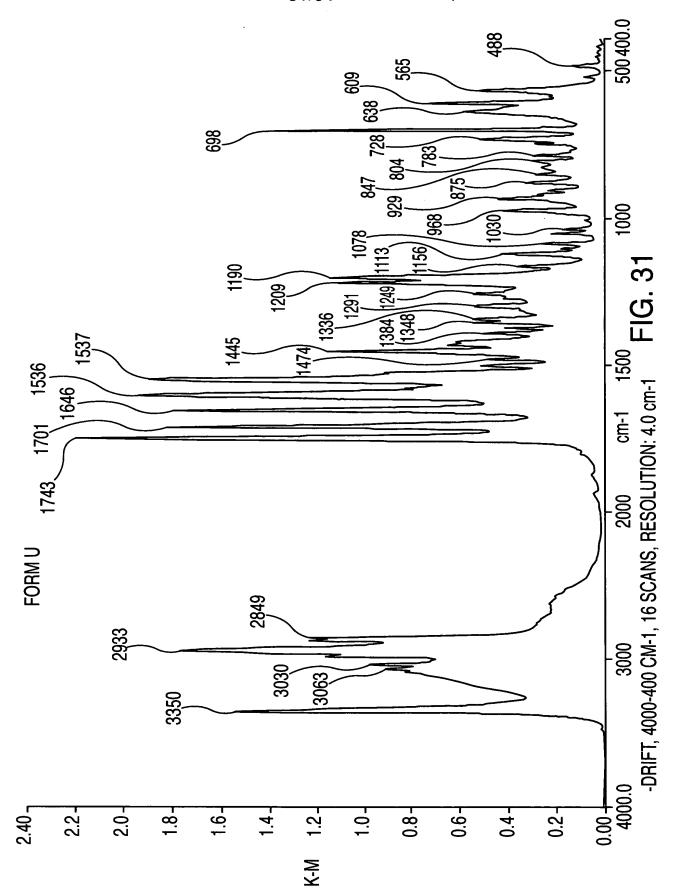
MIXTURE WITH STARTING FORM. WHEN STARTING MATERIAL CONTAINS SEEDS. \*\*\*\*

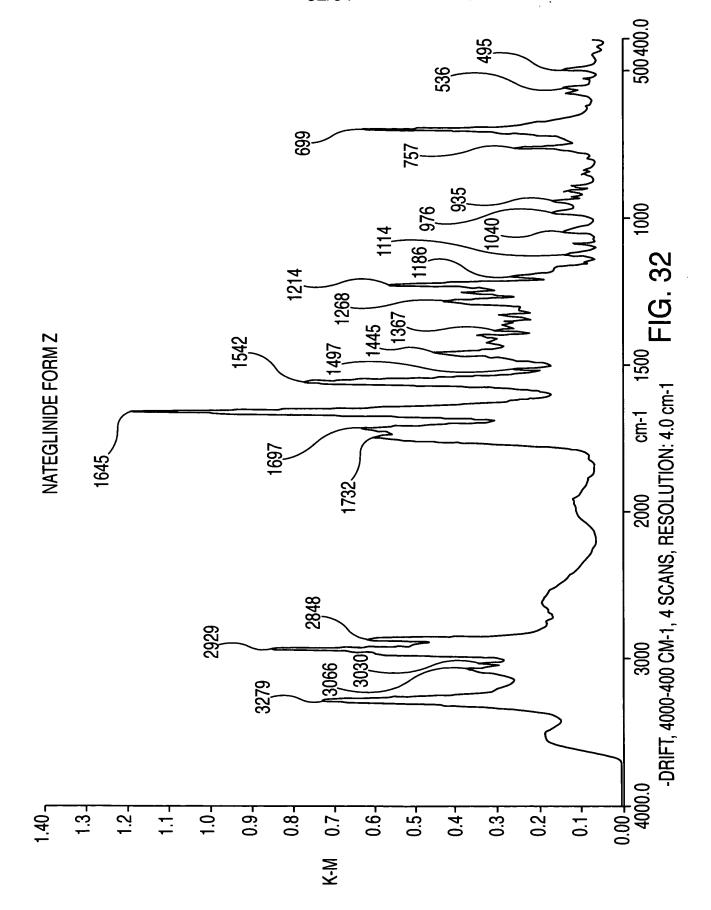
RESULTS MIGHT VARY DEPENDING ON THE SOLVATE OF FORM EPSILON USED.

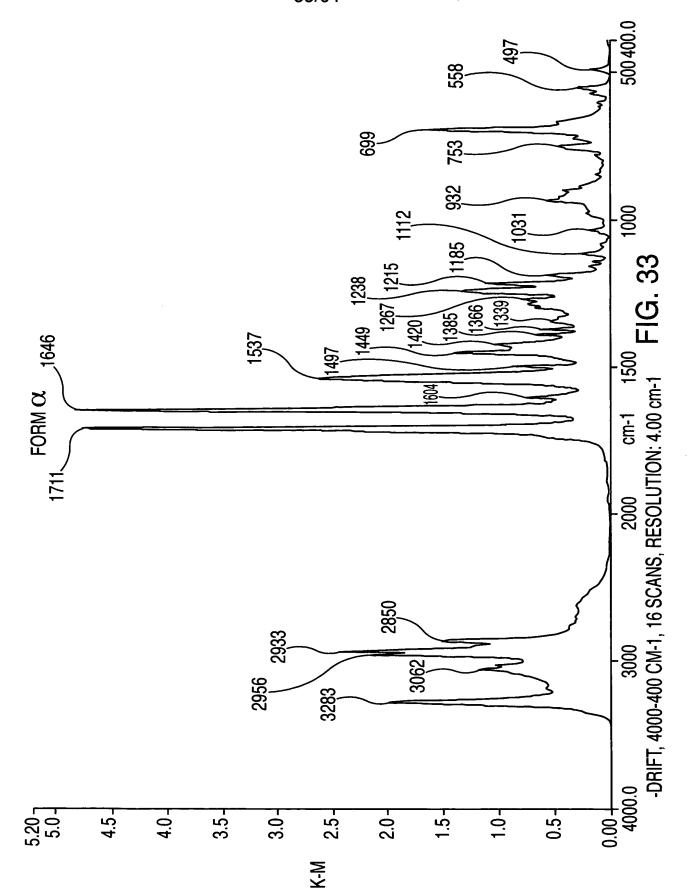
THERMAL STABILITY CHART FIG. 28

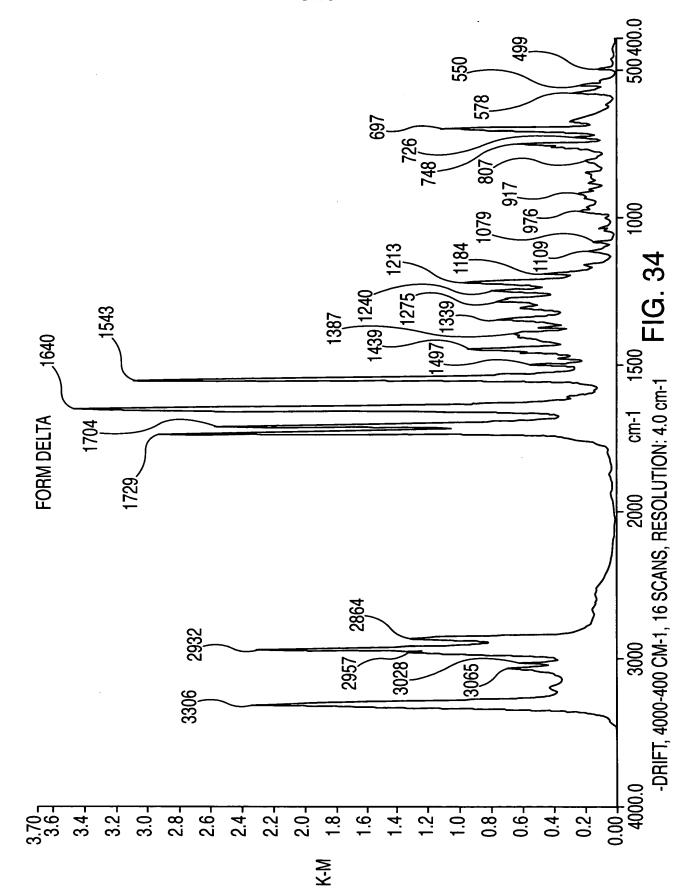


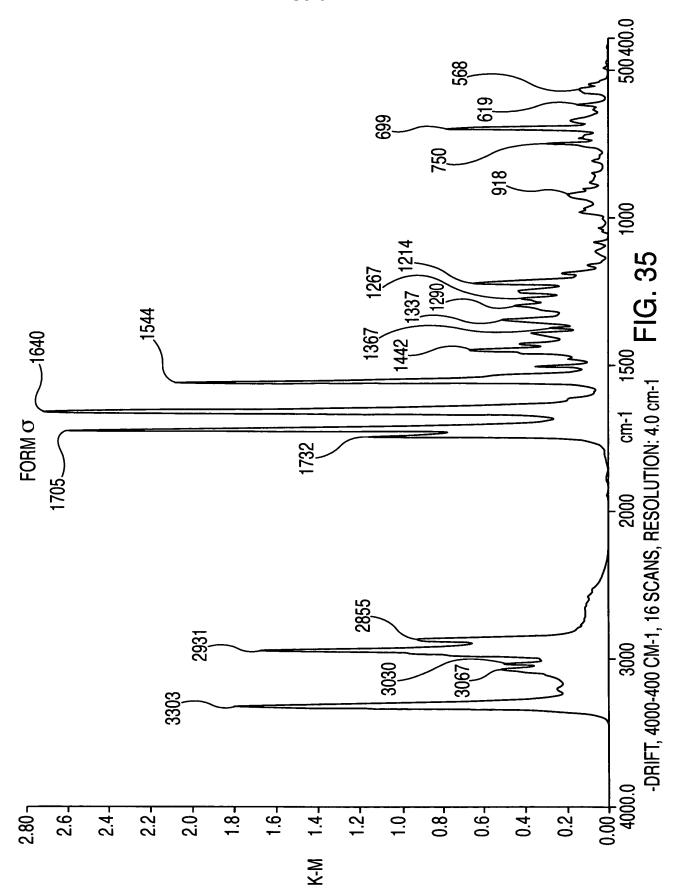


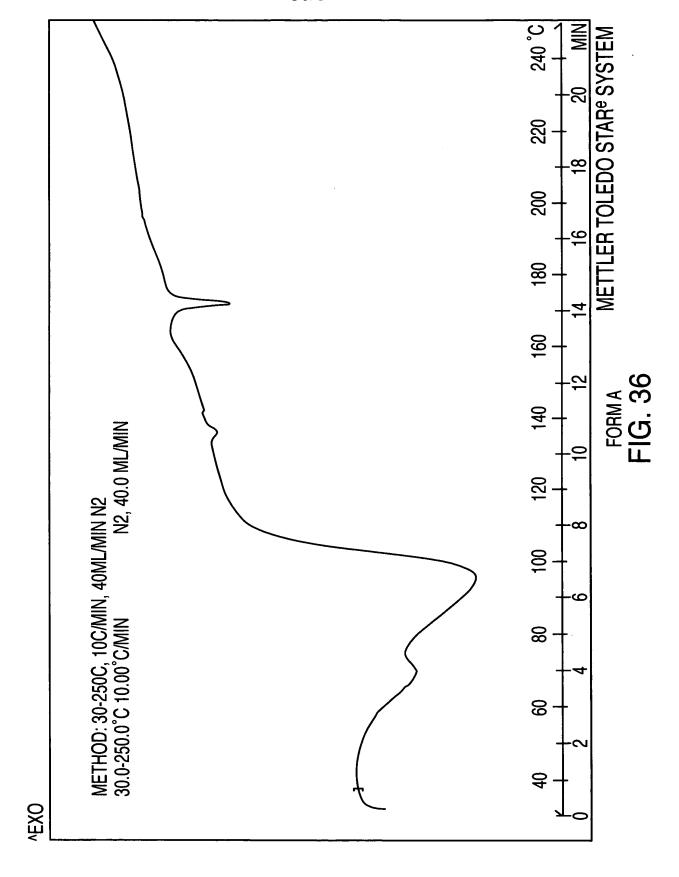


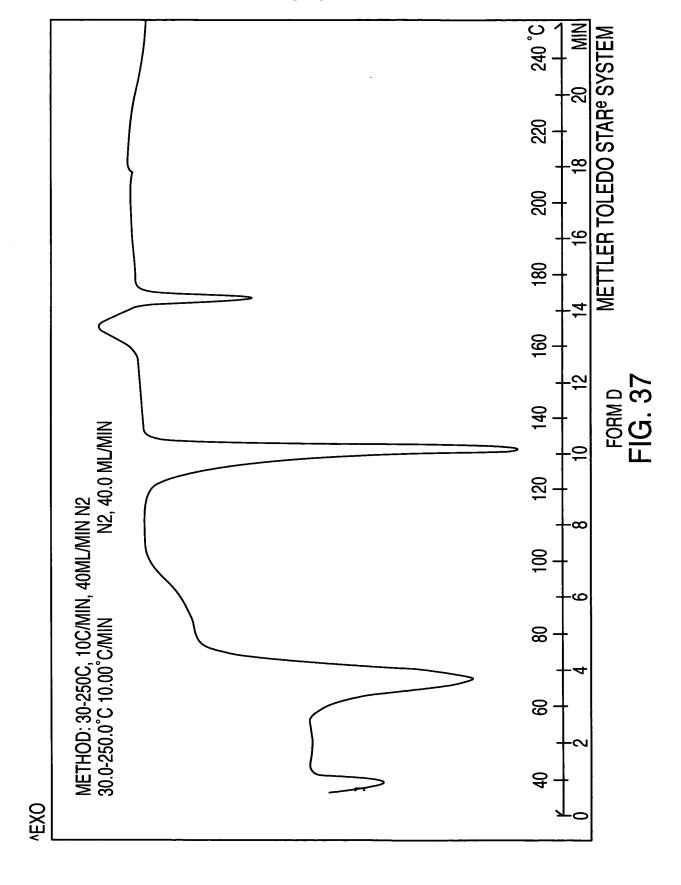


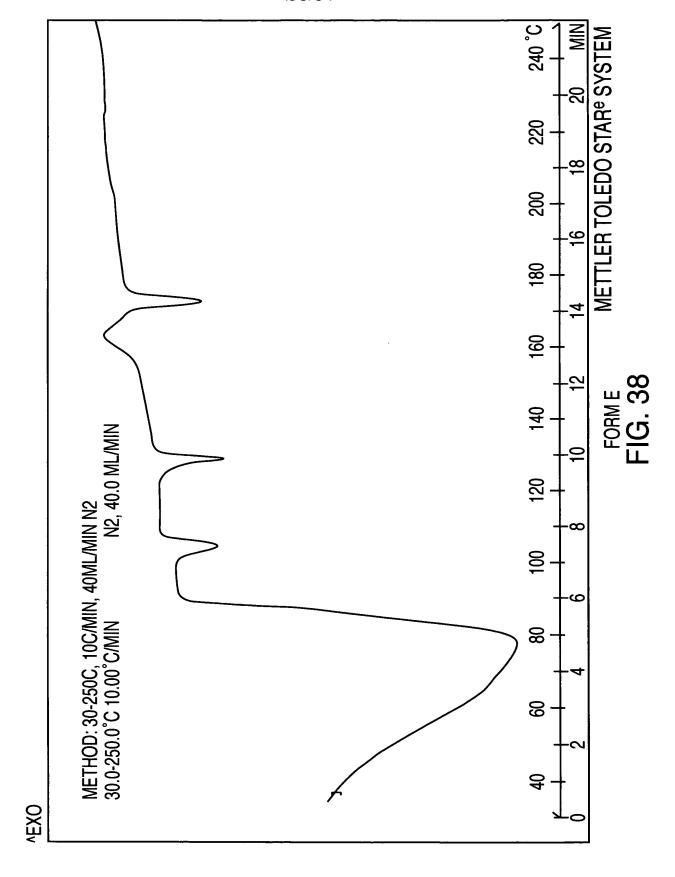


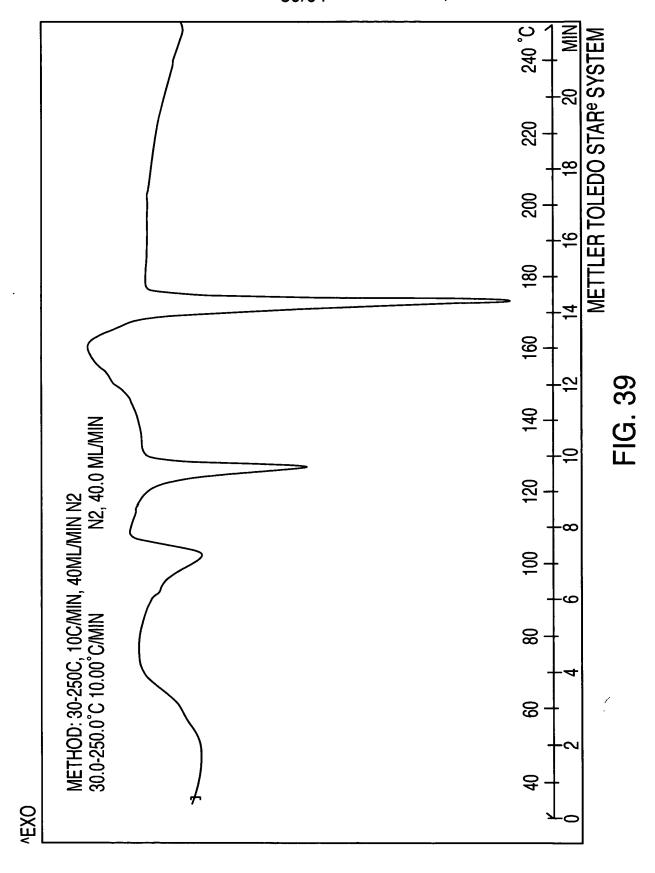


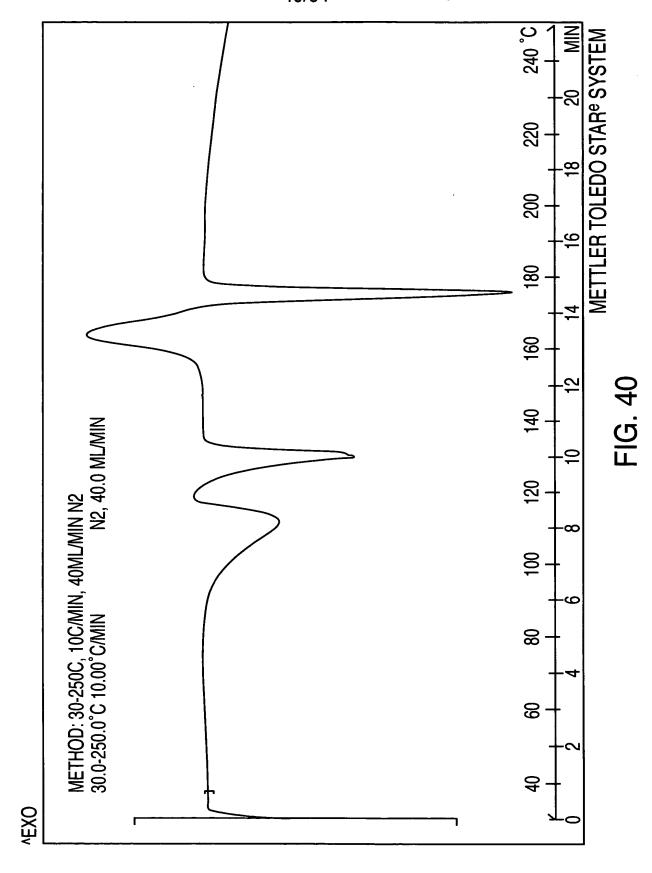


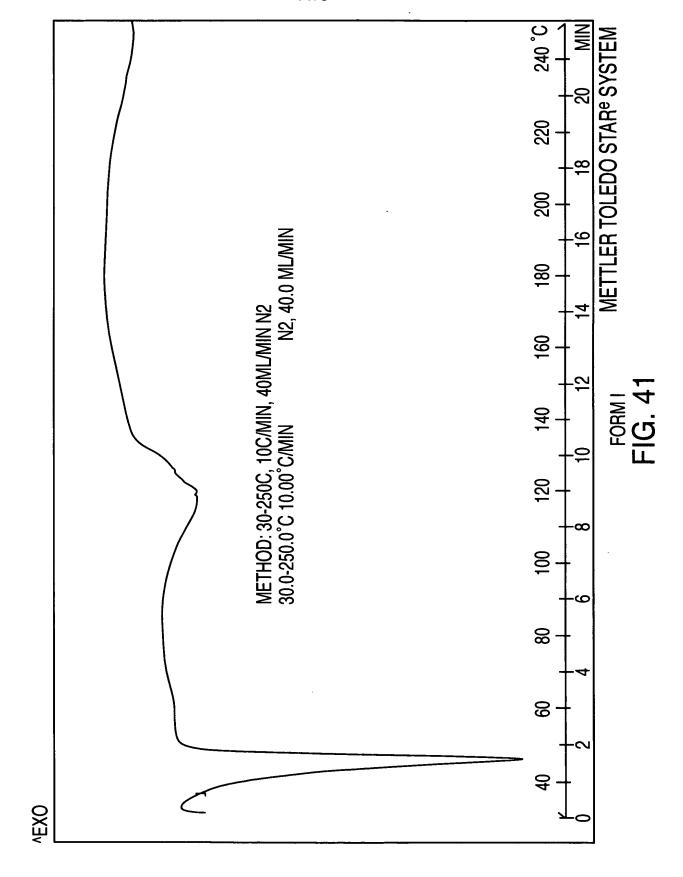


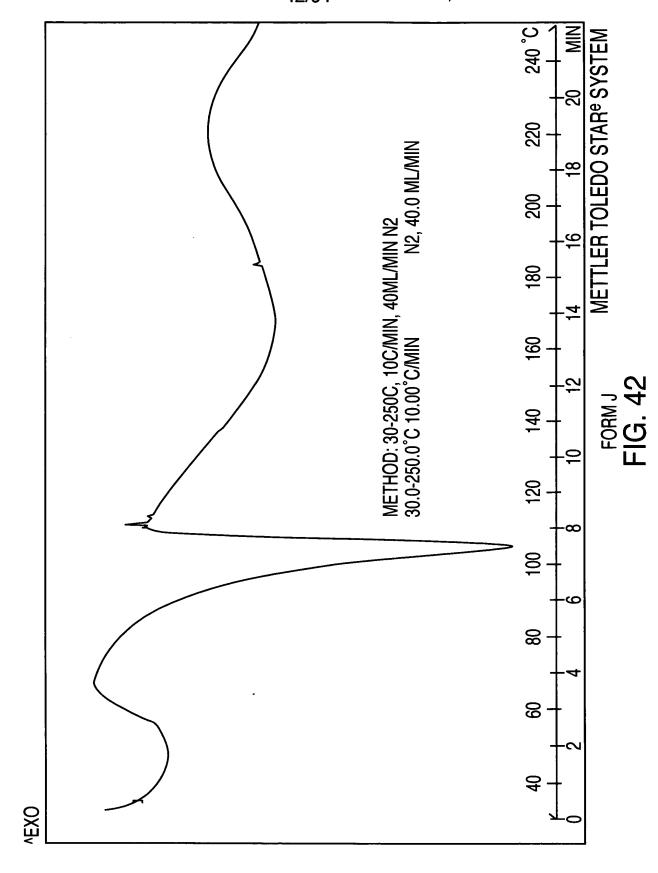


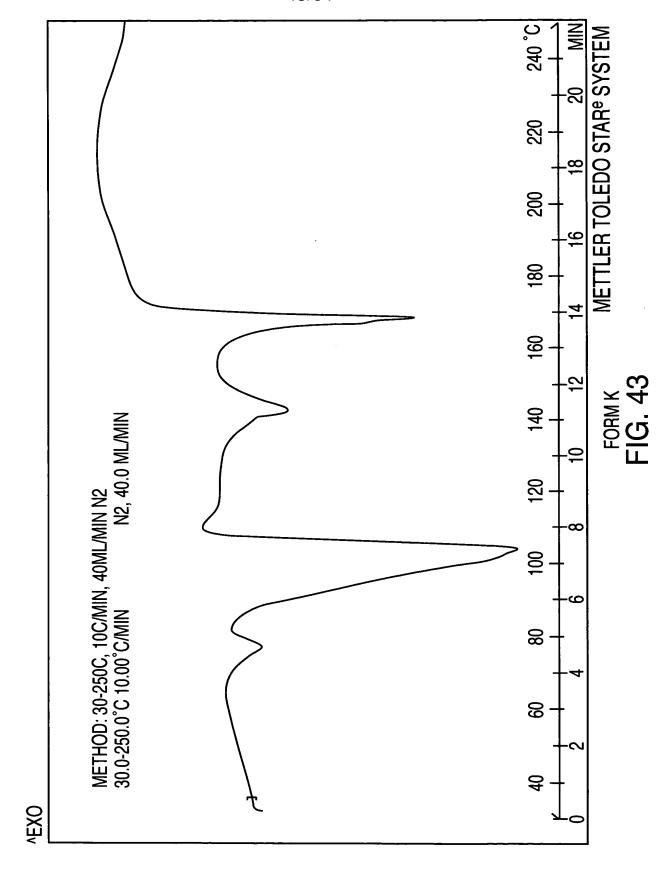


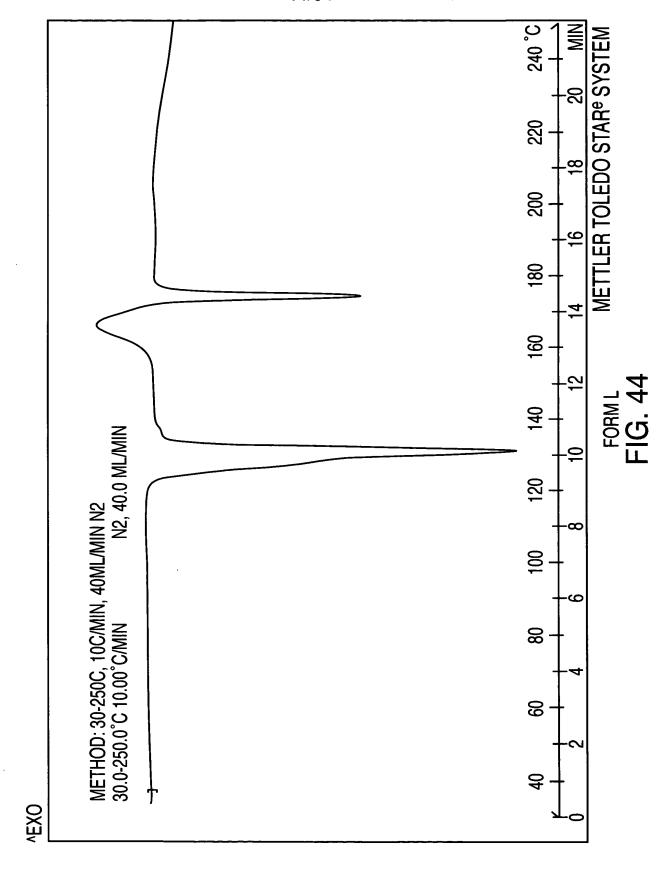


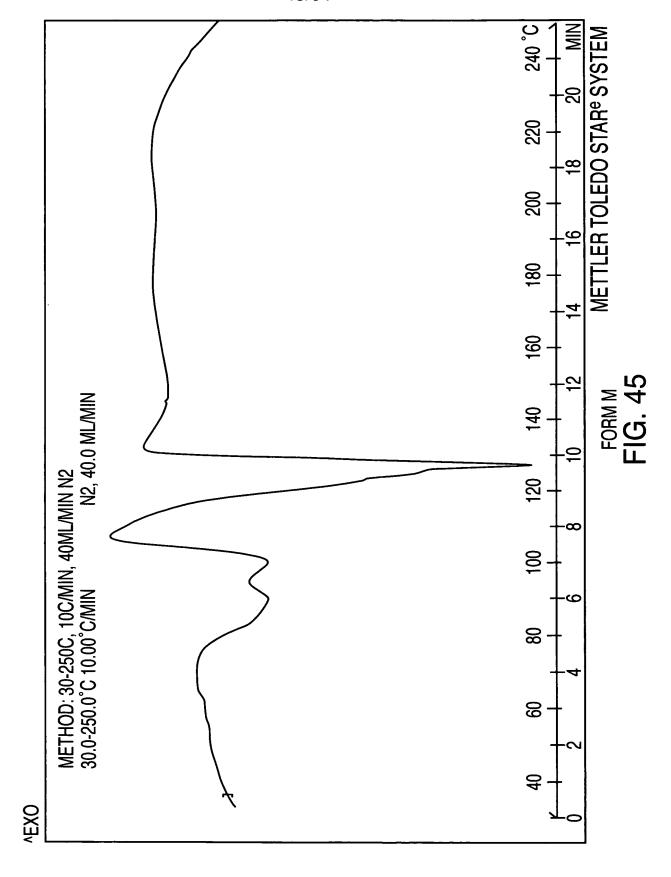


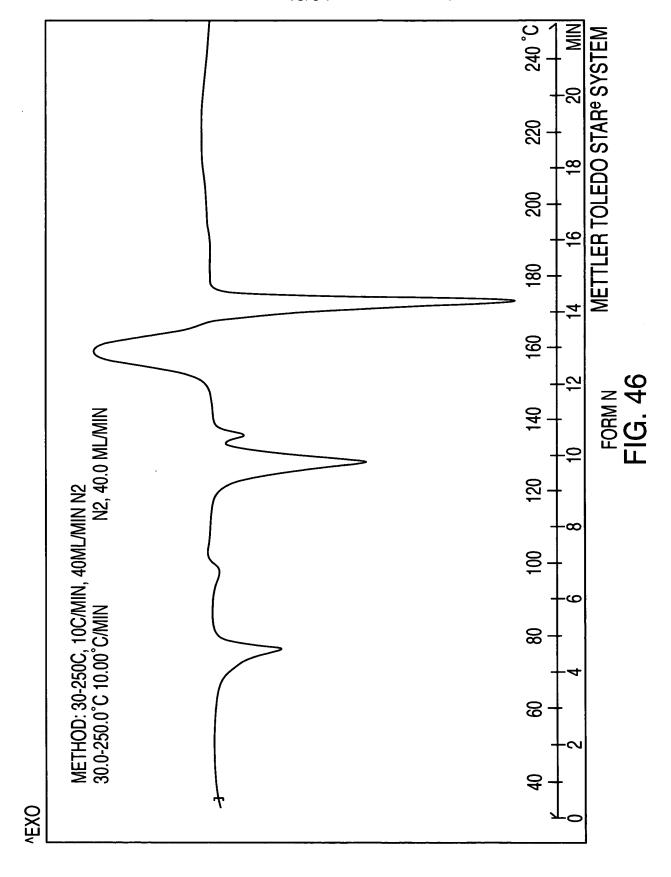


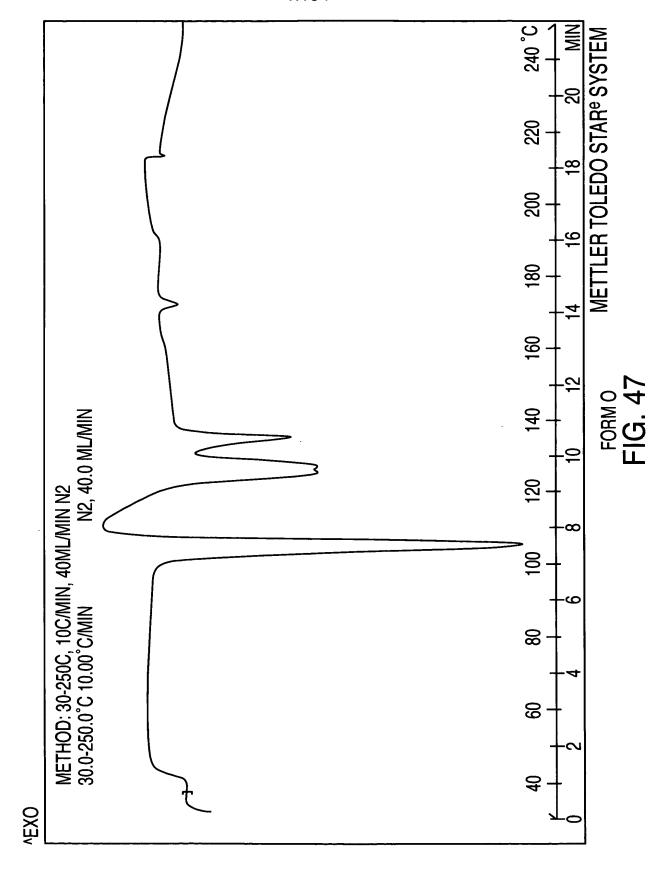


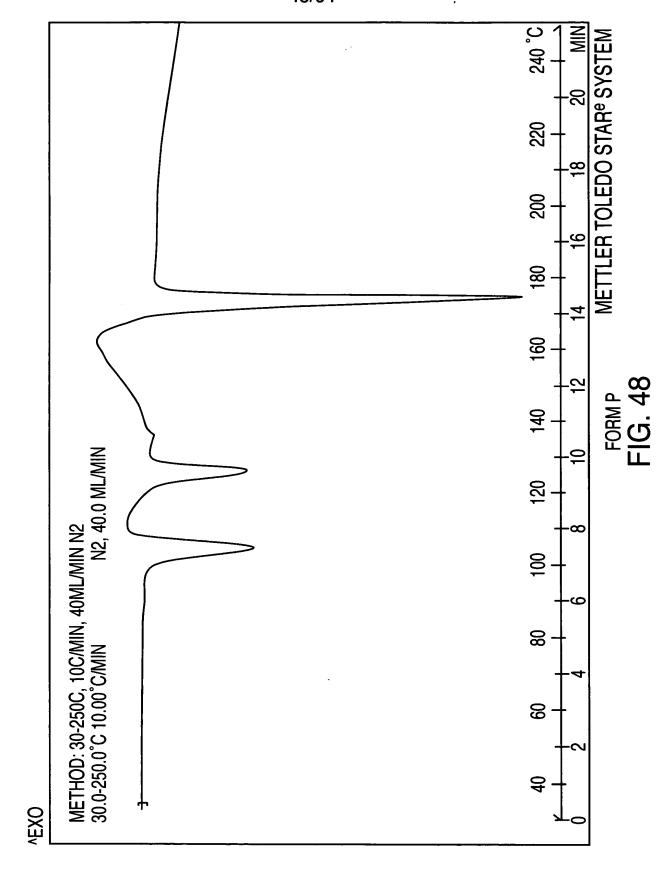


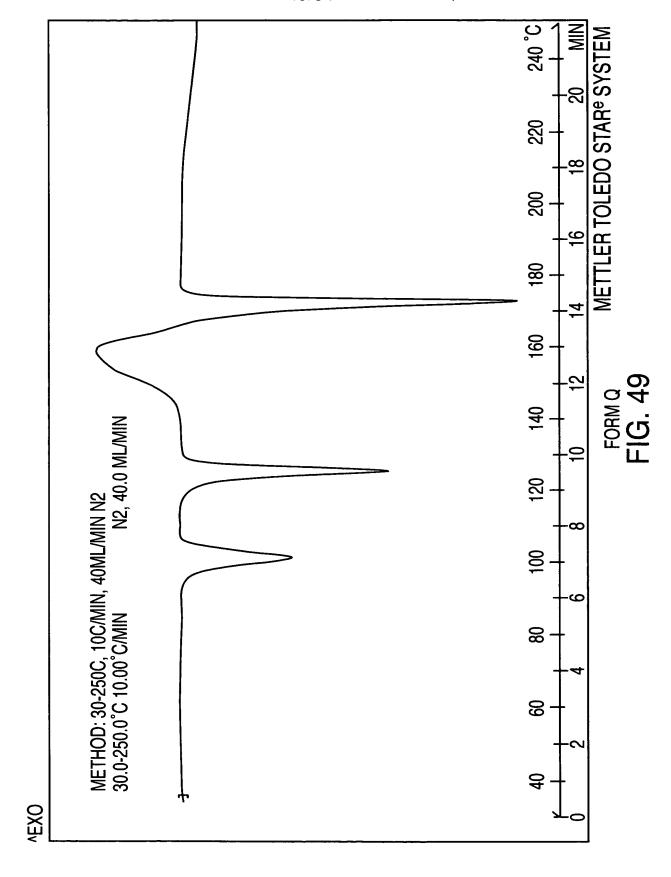


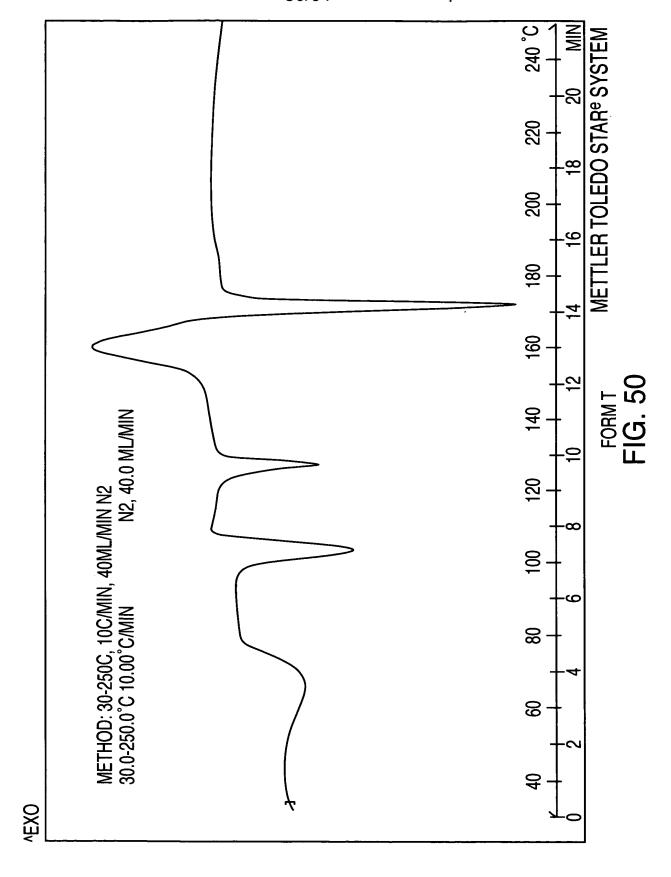


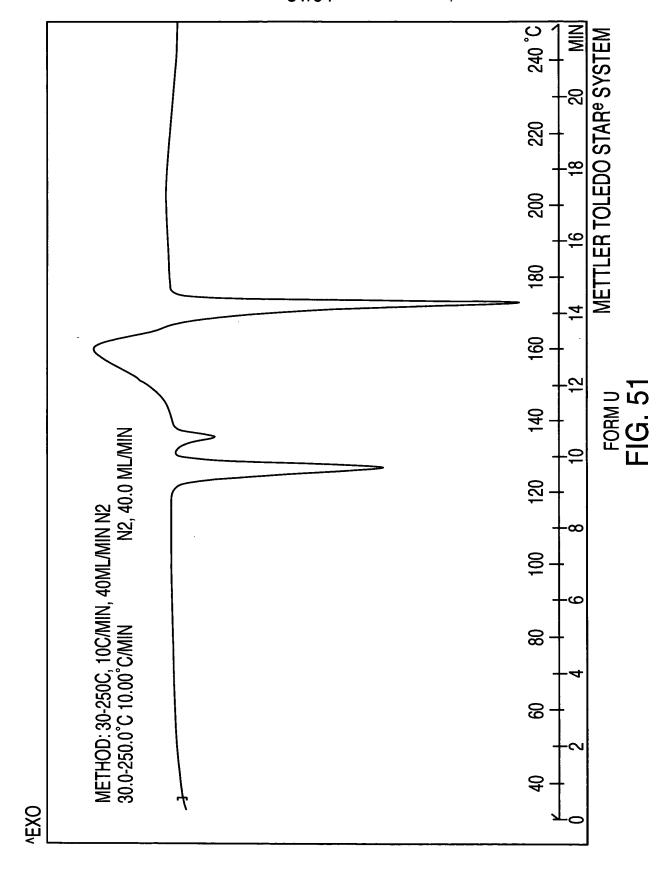


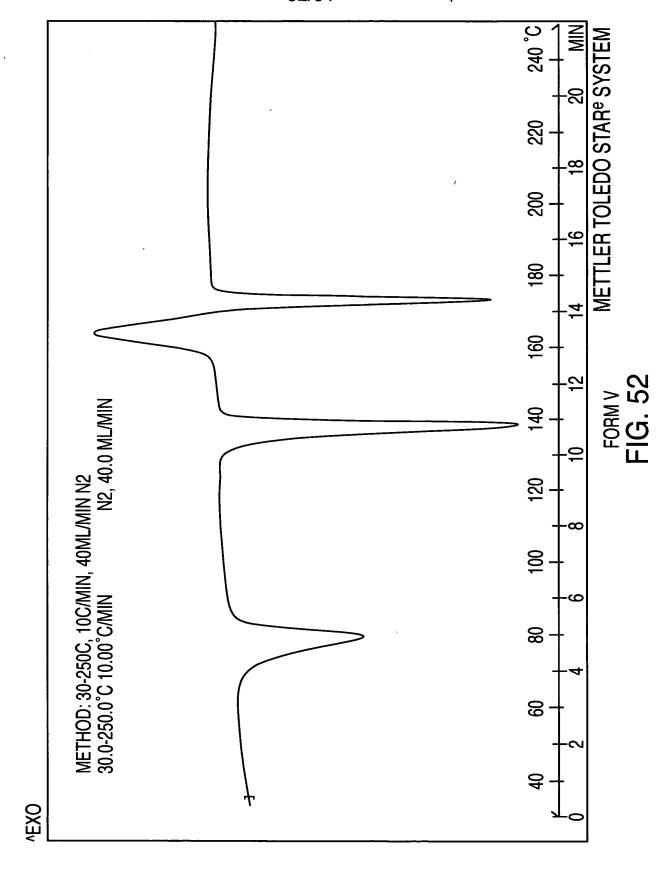


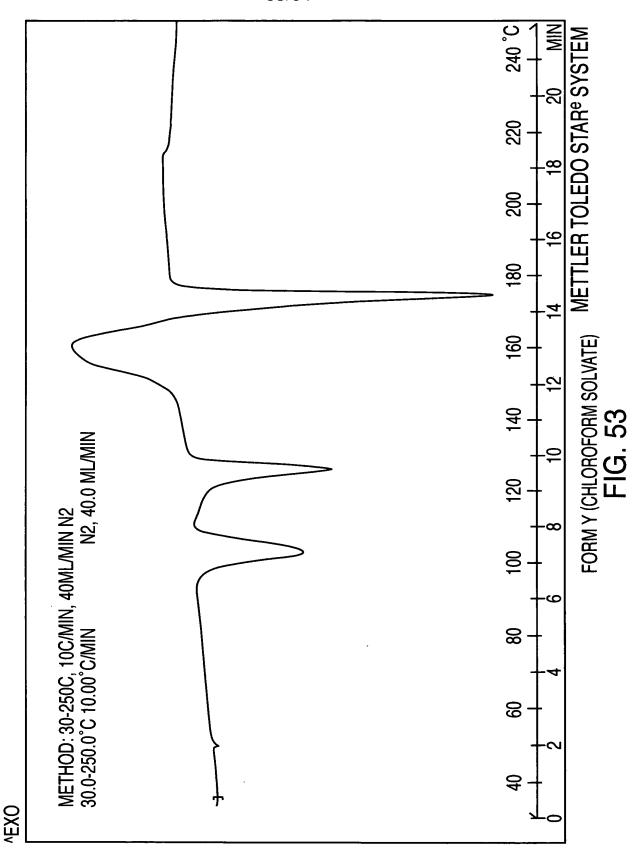


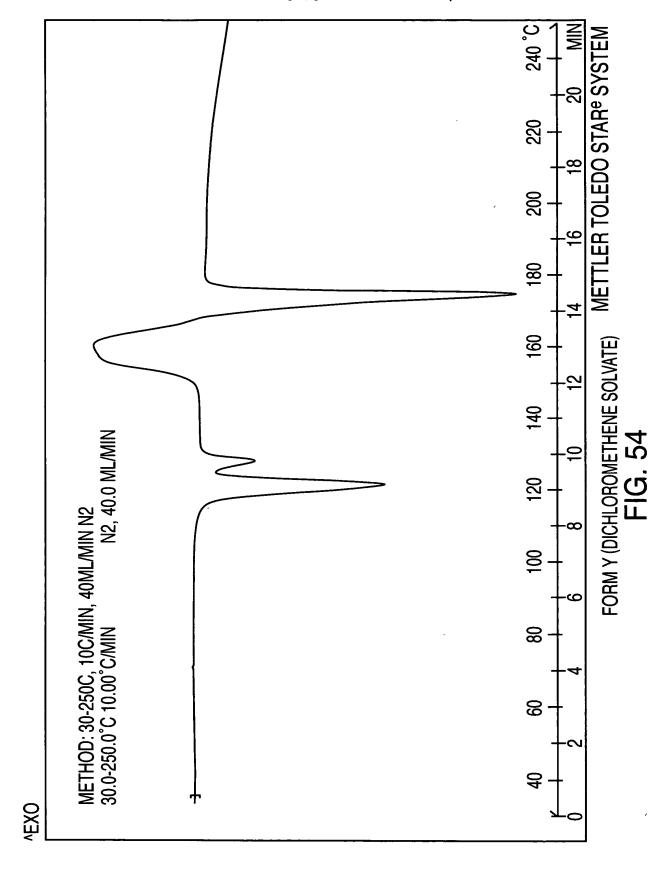


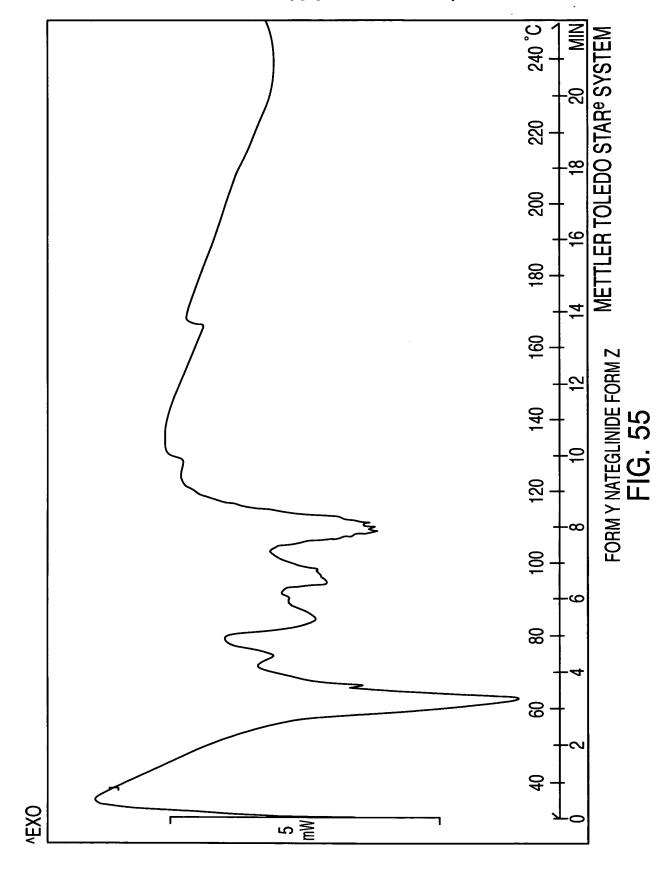


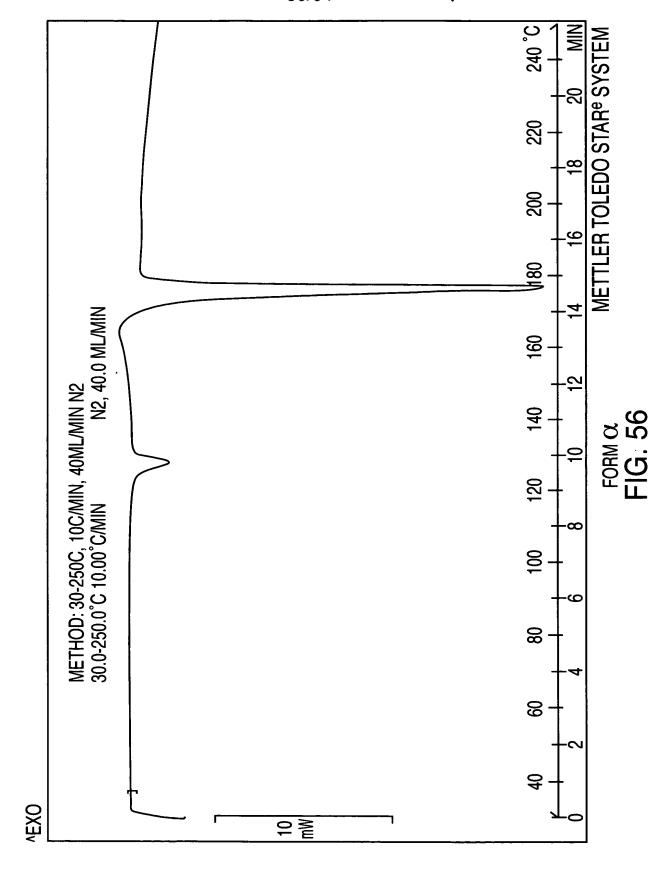


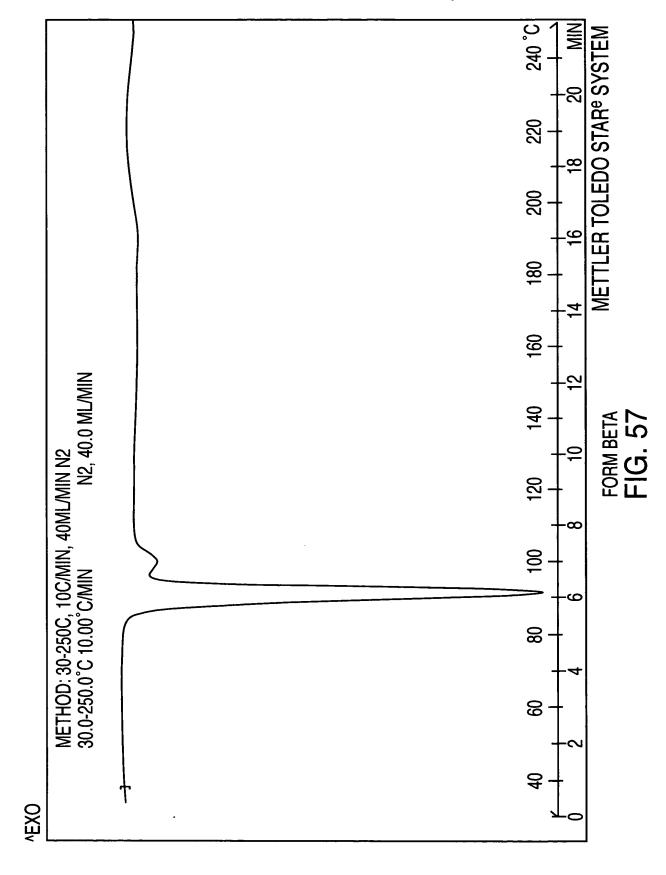


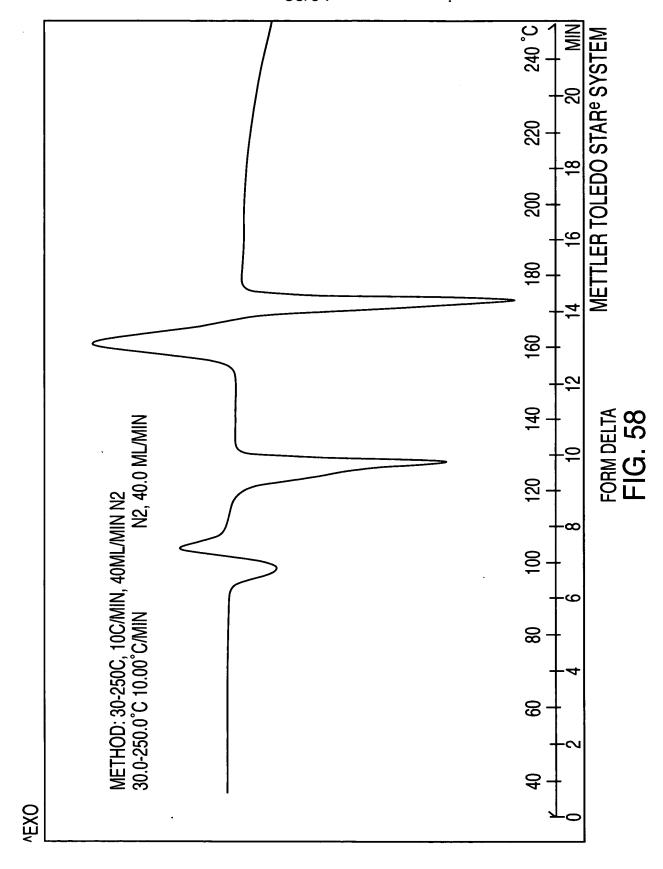


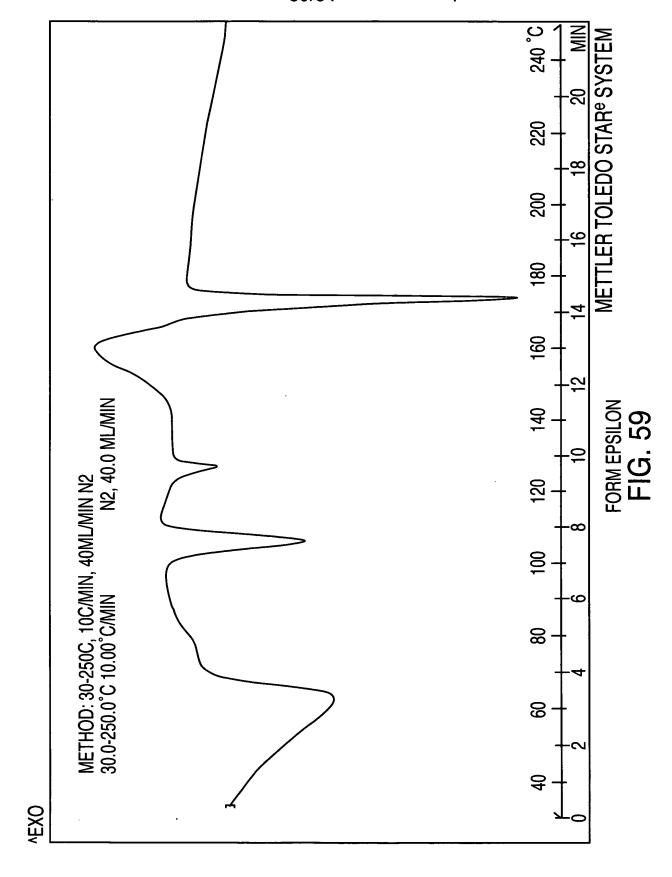


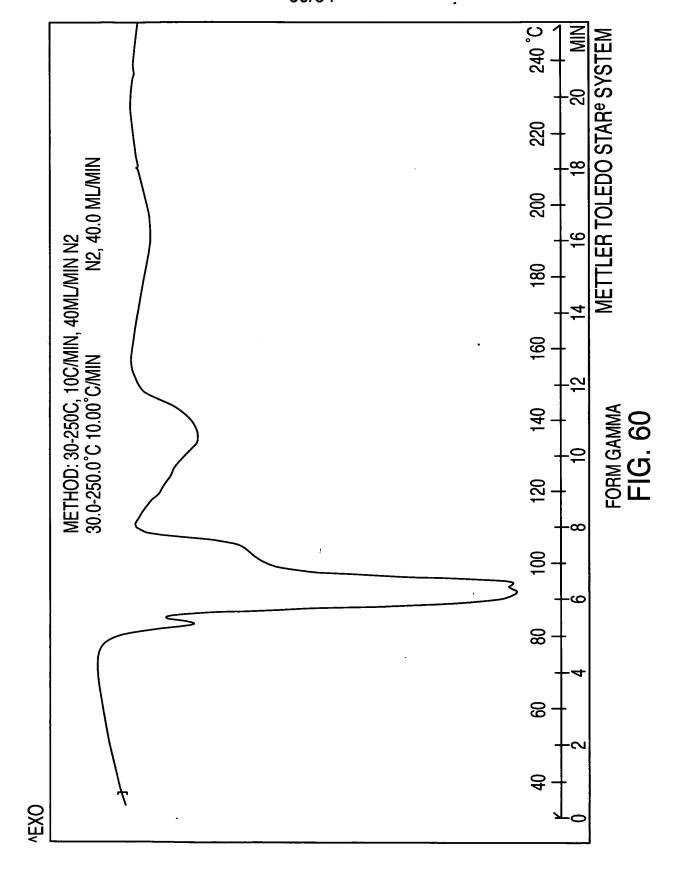


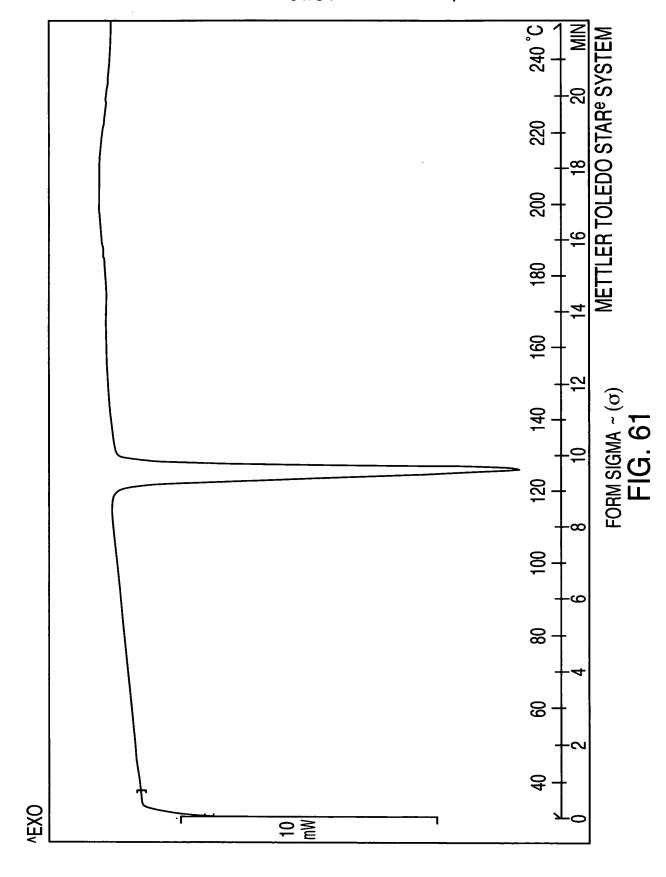


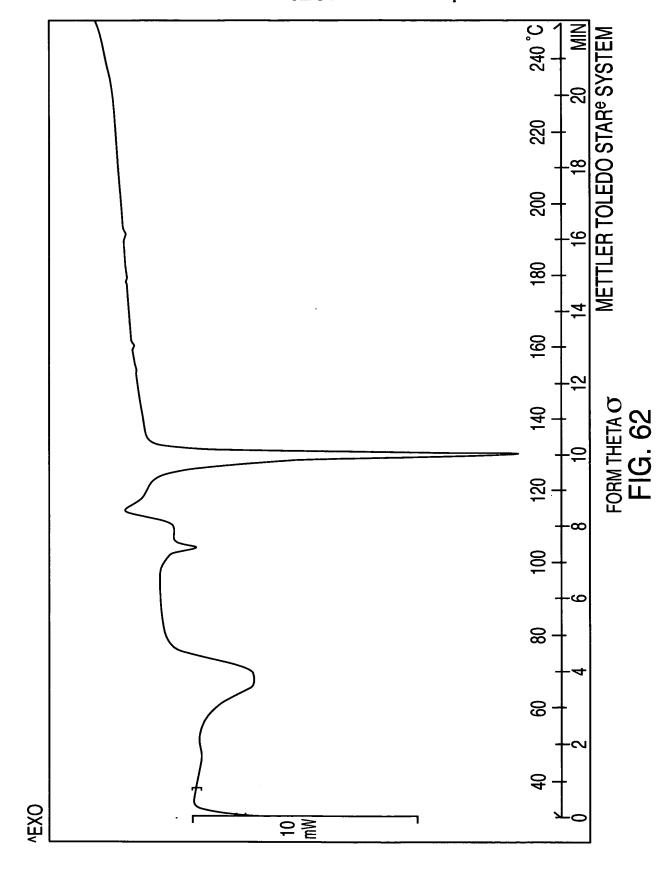


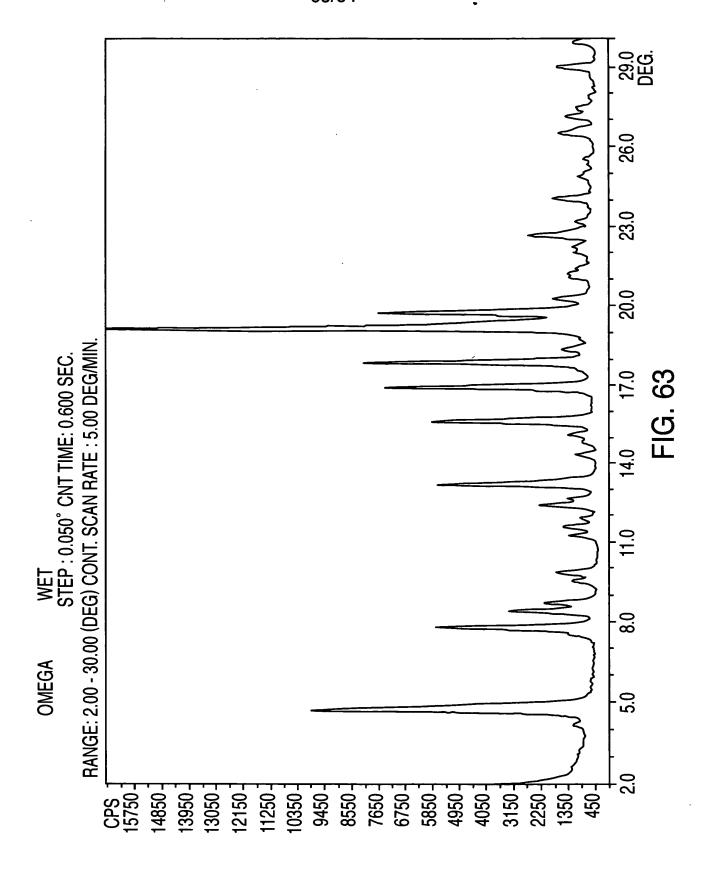












Comparison between the impurity profile of Nateglinide crystallized in IPA-H2O and Nateglinide in Methanol-H2O

Sample No Solvent	Solvent				lm	ourity pi	rfile by F	Impurity prfile by RRT [% w/w]		
		D-PA (0.23)	(0.25)	(0.46)	(0.80)	lpcha (0.89)	Dimer (1.38)	Ipcha Dimer Methyl Ester   (0.89) (1.38) (1.51)	(1.76)	PA (0.25) (0.46) (0.80) Ipcha Dimer Methyl Ester (1.76) Isopropyl Ester (2.3) (0.89) (1.38) (1.51)
RL-2155/1	Methanol-H <sub>2</sub> O <c< td=""><td>&lt;0.01</td><td></td><td>0.02</td><td>&lt;0.01</td><td>0.03</td><td>0.02 &lt;0.01 0.03 0.02</td><td>2.91</td><td>0.04</td><td></td></c<>	<0.01		0.02	<0.01	0.03	0.02 <0.01 0.03 0.02	2.91	0.04	
RL-2163/4	IPA-H <sub>2</sub> 0	<0.01	.01 0.04		0.02	0.02 0.02 0.01	0.01		0.03	0.02

Note: D-PA means D-Phenyl Alanine

Ipcha means Iso propyl cyclohexyl carboxylic acid

Both are the starting materials of the product

(-)-N-[(trans-4-isopropyl cyclohexane)carbonyl]-D-phenylalanine

FIG. 64